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Stroke of a cardiac myxoma origin

Acidente vascular cerebral com origem em mixoma cardíaco

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Abstract

Objective: The clinical features of cardiac myxoma stroke have not been sufficiently described. Debates remain concerning the options and timing of treatment and the clinical outcomes are unknown. This article aims to highlight the pertinent aspects of this rare condition.

Methods: Data source of the present study came from a comprehensive literature collection of cardiac myxoma stroke in PubMed, Google search engine and Highwire Press for the year range 2000-2014.

Results: Young adults, female predominance, single cerebral vessel (mostly the middle cerebral artery), multiple territory involvements and solitary left atrial myxoma constituted the outstanding characteristics of this patient setting. The most common affected cerebral vessel (the middle cerebral artery) and areas (the basal ganglion, cerebellum and parietal and temporal regions) corresponded well to the common manifestations of this patient setting, such as conscious alteration, ataxia, hemiparesis and hemiplegia, aphasia and dysarthria. Initial computed tomography scan carried a higher false negative rate for the diagnosis of cerebral infarction than magnetic resonance imaging did. A delayed surgical resection of cardiac myxoma was associated with an increased risk of potential consequences in particular otherwise arterial embolism. The mortality rate of this patient population was 15.3%.

Conclusion: Cardiac myxoma stroke is rare. Often does it affect young females. For an improved diagnostic accuracy,

magnetic resonance imaging of the brain and echocardiography are imperative for young stroke patients in identifying the cerebral infarct and determining the stroke of a cardiac origin. Immediate thrombolytic therapy may completely resolve the cerebral stroke and improve the neurologic function of the patients. An early surgical resection of cardiac myxoma is recommended in patients with not large territory cerebral infarct.

Descriptors: Embolism. Middle Cerebral Artery. Myxoma. Stroke.

Resumo

Objetivo: As características clínicas do acidente vascular cerebral causado por mixoma cardíaco não foram descritas suficientemente. Debates permanecem sobre as opções e o momento de tratamento e os resultados clínicos são desconhecidos. Este artigo tem como objetivo destacar os aspectos pertinentes desta condição rara.

Métodos: Os dados do presente estudo foram levantados em uma revisão abrangente de literatura sobre acidente vascular cerebral causado por mixoma cardíaco no PubMed, no sistema de buscas do Google e no Highwire Press, abrangendo ao anos entre 2000 e 2014.

Resultados: Adultos jovens, predominância do sexo feminino, vaso cerebral único (principalmente a artéria cerebral mediana), envolvimento de territórios múltiplos e mixoma atrial esquerdo solitário são características marcantes destes pacientes. O vaso

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Abbreviations, acronyms & symbols

CT	Computed tomography
MRI	Magnetic resonance imaging

cerebral afetado mais comum (artéria cerebral média) e áreas (o gânglio basal, cerebelo e regiões parietais e temporais) corresponderam bem com as manifestações comuns destes pacientes, como alteração da consciência, ataxia, hemiparesia e hemiplegia, afasia e disartria. Tomografia computadorizada inicial mostrou taxa de falso negativo mais alta para o diagnóstico de acidente vascular cerebral do que a imagem por ressonância magnética. A ressecção cirúrgica tardia de mixoma cardíaco foi associada com risco aumentado de potenciais consequências, em particular, de outra

INTRODUCTION

Cardioembolic stroke accounts for 14-30% of ischemic strokes with a predilection of early and later recurrences^[1]. Atrial fibrillation, acute myocardial infarction, valvular heart disease, infective endocarditis and cardiac myxoma are major sources of cerebral emboli^[2]. Of them, atrial fibrillation is the most important one responsible for 45% of cardiogenic embolism^[3]. Cardiac myxoma, the most common primary cardiac tumor, is a rare cause, but an important etiology as well for stroke in the young^[4]. However, the diagnosis of a cardiac myxoma is often elusive in young stroke patients^[5]. A delayed diagnosis and untimely treatment may mean *laissez-faire* of progression of stroke and development of certain critical consequences like systemic and peripheral embolic events. The clinical features of stroke of a cardiac myxoma origin have not been sufficiently described. In addition, debates remain concerning the options and timing of treatment and the clinical outcomes are unknown. This article aims to highlight the pertinent aspects of this rare condition.

METHODS

A comprehensive literature collection of cardiac myxoma stroke was made in PubMed, Google search engine and Highwire Press for the year range 2000-2014. The search terms included “cardiac myxoma”, “atrial myxoma”, “valvular myxoma” and “stroke”. The search ended on June 30, 2014. Articles describing transient ischemic attack, cerebral vascular aneurysm without stroke, or stroke caused by other cardiac tumors than myxomas were excluded from this study. Data were extracted from the text or tables, including details of the study subjects, demographics, cerebral infarct, cardiac myxoma, complications, follow-up length and mortality. The main outcomes were complications and mortality.

Quantitative data were expressed in mean \pm standard deviation with range and median values. Comparisons of fre-

forma de embolia arterial. A taxa de mortalidade dessa população de pacientes foi de 15,3%.

Conclusão: Acidente vascular cerebral causado por mixoma cardíaco é raro. Frequentemente, afeta mulheres jovens. Para um diagnóstico mais preciso, exames de ressonância magnética e ecocardiográficos são imperativos para pacientes jovens com acidente vascular cerebral para determinar a localização do enfarte cerebral e se houve origem cardíaca. Terapia trombolítica imediata pode resolver completamente o acidente vascular cerebral e melhorar a função neurológica dos pacientes. Ressecção cirúrgica precoce de mixoma cardíaco é recomendada em pacientes com acidente vascular cerebral de pequena extensão.

Descritores: Embolia. Artéria Cerebral Média. Mixoma. Acidente Vascular Cerebral.

quencies were made by Fisher exact test. P-P plot was used to test normal distribution. Logistic regression was taken to assess the possible predisposing risk factors of mortality. $P < 0.05$ was considered statistically significant.

RESULTS

A total of 83 reports were obtained with 133 patients involved^[4-86]. Eighty-two (98.8%) articles reported only sporadic single case and one (1.2%) article described a small series. There were 71 (53.4%) females and 62 (46.6%) males with a female-to-male ratio of 1.2:1. Patients' ages were 42.3 ± 18.6 (range, 4-84; median, 43.5) years ($n=118$), in a normal distribution. Ages of 67 (56.8%) patients were ≤ 45 years while ages of 51 (43.2%) patients were ≥ 46 years ($\chi^2=4.3$, $P=0.051$). Ages of male and females patients were 40.1 ± 16.1 (range, 10-57; median, 40) years ($n=57$) and 45.0 ± 20.5 (range, 4-84; median, 47) years ($n=64$), respectively ($P=0.093$). The onset time was described in 54 patients with an acute onset in 53 (98.1%) and a chronic onset in 1 (1.9%) patient, respectively ($\chi^2=100.1$, $P=0.000$). Time interval from onset to physician consultation was 1413.1 ± 3581.6 (range, 0.5-17280; median 24) hours ($n=30$) with 17 (56.7%) of them presented within 24 hours, and the remaining 13 patients were admitted 3 days-2 years after the onset.

Their initial symptoms were reported in 108 patients: neurological in 97 (89.8%), constitutional in 3 (2.8%), neurological with constitutional 3 (2.8%), neurological with circulatory in 3 (2.8%), circulatory in 1 (0.9%) and all three triad symptoms in 1 (0.9%) patient. Eleven (8.3%) patients had precursory symptoms including headache in 5 (50%), transient ischemic attack in 2 (20%) and a skin rash/spot in 3 (30%) patients. Totally 104 patients developed neurological symptoms. Hemiparesis, aphasia and conscious alteration were the three most common symptoms (Table 1). Forty-two (31.6%) patients had one or more predisposing risk factors of stroke with hypertension being the most common (Table 2).

Table 1. 228 neurological symptoms in 104 patients.

Neurological symptom	n (%)
Hemiparesis	51 (49.0)
Aphasia	29 (27.9)
Conscious alteration	24 (23.1)
Hemiplegia	18 (17.3)
Dysarthria	17 (16.3)
Ataxia	11 (10.6)
Vision disturbance	10 (9.6)
Dysesthesia	9 (8.7)
Seizures	7 (6.7)
Collapse	6 (5.8)
Dizziness	6 (5.8)
Dysphagia	4 (3.8)
Hemianopsia	4 (3.8)
Mental disturbance	4 (3.8)
Transient ischemic attack	4 (3.8)
Vertigo	4 (3.8)
Diplopia	3 (2.9)
Facial paresis	3 (2.9)
Disorientation	2 (1.9)
Syncope	2 (1.9)
Nausea/vomiting	2 (1.9)
Cognitive impairment	1 (1.0)
Convulsion	1 (1.0)
Epilepsy	1 (1.0)
Hypomnesia	1 (1.0)
Memory loss	1 (1.0)
Paraparesis	1 (1.0)
Ptoxis	1 (1.0)
Quadriparesis	1 (1.0)

Table 2. Predisposing risk factors of stroke.

Risk factor	n (%)
Hypertension	13 (9.8)
Hyperlipidemia	5 (3.8)
Smoking	5 (3.8)
Atrial fibrillation	4 (3.0)
Coronary artery disease	4 (3.0)
Hypertension, hyperlipidemia	3 (2.3)
Hypertension, diabetes mellitus	2 (1.5)
Alcohol	1 (0.8)
Mitral regurgitation	1 (0.8)
Neurosurgery for temporooccipital cavernoma	1 (0.8)
Raynaud's disease	1 (0.8)
Transient ischemic attack	1 (0.8)
Unknown	1 (0.8)

Except for cardiac myxoma and stroke, an additional diagnosis was established in 23 (17.3%) patients including Carney's syndrome in 7 (30.4%), infected cardiac myxoma in 3 (13.0%) (associated with disseminated intravascular coagulation, kidney and spleen infarcts and urinary tract infection in 1 patient each), patent fossa ovalis in 2 (8.7%), and atrial

septal defect (incidental finding during cardiac surgery), patent ductus arteriosus, acute renal impairment, posttraumatic seizure, synovial sarcoma of the right hand, internal carotid artery aneurysm, non-ST segment elevation myocardial infarction, pregnancy, Raynaud's disease plus multiple cerebral aneurysms, NAME syndrome and systemic vasculitis with antiphospholipid antibody syndrome in 1 (4.3%) patient each.

Erythrocyte sedimentation rate was detected in 25 (18.8%) patients: 23 (92%) were positive and 2 (8%) were negative ($\chi^2=35.3$; $P=0.000$) with a quantitative value of 57.4 ± 19.7 (range, 30-85; median, 60) mm/h ($n=15$). C-reaction protein was detected in 16 (12.0%) patients: positive in 10 (62.5%) and negative in 6 (37.5%) patients ($\chi^2=2.0$; $P=0.289$) with a quantitative value of 8.5 ± 13.0 (range, 0.09-35; median, 1.8) mg/dL ($n=10$).

Cardiac myxoma was diagnosed ahead of the diagnosis of stroke in 8 (11.8%), delayed until after the diagnosis of stroke in 59 (86.8%) and simultaneously with the diagnosis of stroke in 1 (1.5%) patient.

The diagnostic techniques for stroke were given in 81 (60.9%) patients with magnetic resonance imaging (MRI) being the most commonly and computed tomography (CT) more commonly used technique (Table 3).

Table 3. Diagnostic tools for cerebral infarct in 81 patients.

Diagnostic tool	n (%)
MRI	39 (48.1)
CT	23 (28.4)
CT, MRI	7 (8.6)
MRI, MRA	4 (4.9)
MRA	3 (3.7)
Autopsy	1 (1.2)
CT, MRA	1 (1.2)
CT, MRI, angiogram	1 (1.2)
Computed tomographic angiography	1 (1.2)
MRI, autopsy	1 (1.2)

CT=computed tomography; MRI=magnetic resonance imaging; MRA=magnetic resonance angiography.

Table 4. Results of computed tomography and time of scanning, n (%).

Computed tomography	Initial	Subsequent
Positive	5 (26.3)	40 (87.0)
Negative	14 (73.7)	6 (13.0)
χ^2	8.5	50.3
P value	0.009	0.000

Sixty-five CT scans were examined in 53 patients with a false negative rate of 30.8% (20/65) in overall and of 73.7% (14/19) in initial CT scans taken within 3 hours after the stroke onset (Table 4). Sixty MRIs were taken in 58 patients: 58 (96.7%) MRIs illustrated a positive result and only 2 (3.3%) were negative, both of which were taken within 7 hours after

the onset. The false negative rate of MRI was much lower than that of CT scan ($\chi^2=15.2$, $P=0.000$). Magnetic resonance angiography was investigated in 14 (10.5%) patients: positive in 12 (85.7%) and unremarkable in 2 (14.3%) patients ($\chi^2=14.3$, $P=0.000$). Cerebral angiogram was performed in only one patient 3 hours after the onset and it revealed a normal result.

The affected cerebral artery was reported in 54 patients and the infarct region could be tracked in 62 patients. The most common affected vessel was the middle cerebral artery (Table 5) and the most common infarct regions were the basal

Table 5. The affected cerebral artery in 54 patients.

Affected cerebral artery	n (%)
MCA	35 (64.8)
Left	21 (60)
Right	12 (34.3)
Unknown	2 (5.7)
Internal carotid artery	6 (11.1)
Posterior cerebral artery	3 (5.6) [1 (33.3) with aneurysm and 1 (33.3) with stenosis]
Basilar artery	2 (3.7)
MCA, anterior cerebral artery	2 (3.7)
Internal carotid artery + vertebrobasal artery	1 (1.9)
MCA, posterior inferior cerebellar artery	1 (1.9)
MCA, posterior cerebral artery	1 (1.9)
MCA, basilar artery	1 (1.9)
Posterior inferior cerebellar artery	1 (1.9)
Superior cerebellar artery	1 (1.9)

MCA=middle cerebral artery.

Table 6. Locations of cerebral infarcts in 62 patients.

Location of infarct	n (%)
Basal ganglion	28 (45.2)
Cerebellum	23 (37.1)
Parietal	17 (27.4)
Temporal	12 (19.4)
Cerebral	10 (16.1)
Frontal	9 (14.5)
Occipital	9 (14.5)
Capsula interna	7 (11.3)
Thalamic	6 (9.7)
Frontoparietal	5 (8.1)
Brainstem	3 (4.8)
Lacunar	3 (4.8)
Pons	3 (4.8)
Corpus callosum	2 (3.2)
Corona radiata	1 (1.6)
Frontotemporal	1 (1.6)
Insular cortex	1 (1.6)
Medulla	1 (1.6)
Perisylvian	1 (1.6)
Periventricular	1 (1.6)
Sylvian fissure	1 (1.6)
Watershed	1 (1.6)

ganglion, cerebellum and parietal and temporal regions (Table 6). An old cerebral infarct was noted in 6 (4.5%) patients.

Patients were more with unilateral cerebral infarct than with bilateral ($\chi^2=14.1$, $P=0.000$), and more with multiple infarcts than solitary. No prevalence was noted between left- and right-sided infarcts (Table 7).

Nineteen (14.3%) patients had other than cerebral artery embolisms, with systemic multiple emboli in 4 (21.1%)^[55,63,71,80], peripheral multiple emboli in 3 (15.8%)^[19,72,85] and solitary arterial embolus in 12 (63.2%) patients^[9,11,13,16,21,35,41,43,60,76,84]. The 39 affected arteries in 18 patients were listed in Table 8.

Table 7. Comparisons of site and number of cerebral infarcts.

Infarct side	Single	Multiple	χ^2	P value
Left	15 (39.5)	14 (22.6)	0.069	1.000
Right	14 (36.8)	6 (9.7)	6.40	0.026
Bilateral	--	26 (41.9)	--	--
Not given	9 (23.7)	16 (25.8)	3.92	0.089
Total	38 (100)	62 (100)	11.52	0.001

Table 8. The 39 affected arteries other than cerebral artery in 18 patients.

Affected artery	n (%)	Reference
Extremity artery	14 (35.9)	
Brachial	1 (7.1)	[72]
Radial	1 (7.1)	[72]
Ulnar	1 (7.1)	[72]
Dorsalis pedis	1 (7.1)	[85]
Peroneal	1 (7.1)	[72]
Popliteal	2 (14.2)	[80]
Tibial	2 (14.2)	[72,85]
Tibio-peroneal trunk	1 (7.1)	[72]
Unknown	4 (28.6)	[13,76,84]
Visceral artery	13 (23.1)	
Hepatic	1 (7.7)	[63]
Mesenteric	2 (15.4)	[16,80]
Renal	3 (23.1)	[55,63,80]
Splenic	3 (23.1)	[55,63,80]
Coronary	4 (10.3)	
Circumflex	1 (25)	[41]
Ramus intermedius	1 (25)	[35]
Unknown	2 (50)	[63,80]
Other peripheral artery	8 (20.5)	
Carotid	3 (37.5)	[11,19,63]
Femoral	2 (25)	[21,80]
Iliac, internal	2 (25)	[55,80]
Retinal	1 (12.5)	[9]
Great vessel	4 (10.3)	
Abdominal aorta	2 (50)	[60,80]
Infrarenal aorta	1 (25)	[19]
Pulmonary	1 (25)	[43]

On admission, an immediate thrombolytic therapy with recombinant tissue plasminogen activator (rtPA) was given in 9 (6.8%) patients with an onset-to-treatment interval of 104.2 ± 36.9 (range, 65-160; median, 92) minutes ($n=5$). The thrombolytic therapy was successful in 7 (77.8%), failed in 1 (11.1%) and the effect unavailable in 1 (11.1%) patient. However, one of the successful patients died of diffuse cerebral edema in spite of an urgent decompressive craniectomy^[21]. On admission, the Glasgow Coma Scale of the patients was 10 ± 2.6 (range, 7-15; median, 9) ($n=13$) and NIH Stroke Scale was 16.7 ± 5.8 (range, 10-26; median, 15.5) ($n=10$). The latter was decreased to 9.7 ± 7.6 (range, 0-21; median, 9) ($n=6$) after treatment ($P=0.055$).

Timing of the diagnosis of cardiac myxoma was described in 33 (24.8%) patients. A patient had been diagnosed with a cardiac myxoma 7 years earlier during which time she declined to all treatments. The remaining 32 (97.0%) patients had their cardiac myxomas diagnosed at a mean of 342.7 ± 744.6 (range, 1-3240; median, 7.5) days after admission.

The diagnostic techniques of cardiac myxoma were described in 114 (85.7%) patients. Transthoracic echocardiography was the most common diagnostic technique, which was used in 90 (78.9%) patients. It was the only diagnostic tool in 79 (87.8%) patients and as an adjunctive to other imaging in 11 (12.2%) patients (Table 9). In one patient, the pathology of the aspirated materials by catheterized thrombectomy on day 5 after admission revealed platelet thrombus and myxomatous tissue, which aroused the suspicion of a cardiac myxoma origin^[41]. Both transthoracic and transesophageal echocardiography showed a respective false negative result in one patient each^[10,19], the cause of the latter was a complete detachment of the cardiac myxoma^[19].

The left atrium was the most common location of cardiac myxoma. There were more solitary myxomas than multiple (Table 10). The attachments of myxomas were reported in 32 myxomas of 31 (23.3%) patients: 25 (78.1%) myxomas were pedunculated and 7 (21.9%) were sessile ($\chi^2=20.3$, $P=0.000$).

Table 9. Diagnostic means of cardiac myxoma.

Diagnostic means	n (%)
TTE	79 (69.3)
TTE, TEE, 3D TEE	12 (10.5)
TEE	9 (7.9)
TTE, MRI	4 (3.5)
Autopsy	3 (2.6)
Computed tomography, TTE	2 (1.8)
TTE, TEE, MRI	2 (1.8)
Computed tomography, TTE, TEE	1 (0.9)
Computed tomography angiography, TTE	1 (0.9)

MRI=magnetic resonance imaging; TEE=transesophageal echocardiography; TTE= transthoracic echocardiography

Prolapse of atrial myxoma into the ventricle was described in 45 (33.8%) patients with a complete prolapse in 43 (95.6%), a partial prolapse in 1 (2.2%) and no prolapse in 1 (2.2%) patient. Gross appearance of the myxoma was reported in 52 (39.1%) patients: irregular in 48 (92.3%) and smooth in 4 (7.7%) ($\chi^2=74.5$, $P=0.000$). The tumor size was 40.0 ± 19.7 (range, 2-92; median, 40.4) mm ($n=71$).

Table 10. Locations of 112 cardiac myxomas.

Location	n (%)
Single	105 (93.8)
LA	93 (88.6)
LV	8 (7.6)
Left heart	1 (1.0)
Atrium	1 (1.0)
Infrarenal aorta (complete detachment to)	1 (1.0)
RA	1 (1.0)
Multiple	7 (6.3)
LA, RA	2 (28.6)
LA, LV, RA, RV	1 (14.3)
Multiple chambers	1 (14.3)
LA, pulmonary vein orifice, mitral annulus	1 (14.3)
LA, mitral annulus	1 (14.3)
LA, RV	1 (14.3)

LA=left atrium; LV=left ventricle; RA=right atrium; RV=right ventricle.

Timing of cardiac myxoma resection was reported in 48 (36.1%) patients. Seven (14.6%) patients underwent operation urgently without giving an exact time interval from admission to surgery. In the remaining 41 (85.4%) patients, the interval from admission to surgery was 140.5 ± 439.9 (range, 1-2550; median, 21) days. The causes of a delayed operation in 3 patients were requirement of heparin therapy for 1 week^[61], minimizing the risks from cardiopulmonary bypass and heparinization^[25] and avoidance of exacerbation of cerebral hemorrhage^[33] in one patient each, and the operations were delayed for 7, 21 and 28 days, respectively. The interval between the diagnosis and surgical operation of cardiac myxoma was 194.2 ± 798.5 (range, 0-4302; median, 12) days ($n=38$). One (2.6%) patient was operated on immediately, 2 (5.3%) patients, within 24 hours, and 14 (36.8%) patients, within one week. A delayed myxoma resection was performed in 42.1% (8/19) patients with alternative embolic events in comparison to 25.4% (29/114) patients without alternative embolic events. The interval from diagnosis to treatment of the former was significantly longer than that of the latter (872.5 ± 1644.2 days vs. 13.3 ± 11.2 days, $P=0.005$).

The treatment methods were unavailable in 28 (21.1%) patients. Nine (6.8%) patients did not receive a cardiac myxoma resection due to sudden death in 3 (33.3%)^[18,60,67], patient decline in 2 (22.2%)^[4,35], and poor patient condition only indicated for anticoagulant therapy^[40], rapid deterioration in spite

of peripheral embolectomies and herniation decompressive hemicraniectomy^[19], unfit for cardiac surgery but only cranial decompression^[63] and extensive metastasis in the lungs^[73] in 1 (12.5%) patient each. The surgical operations performed in 98 (73.7%) patients, 82 (83.7%) of which were isolated cardiac myxoma resection and 2 (2.0%) of which received an operation other than cardiac myxoma resection, were listed in Table 11. Patients without a myxoma resection showed an increased mortality (44.4%, 4/9) than those receiving a myxoma resection (2.1%, 2/95) ($\chi^2=27.1$, $P=0.000$). Of the 9 patients receiving a myxoma resection on an urgent basis, 1 patient died with a mortality of 11.1%, while no patient died among the 45 patients with a delayed surgical resection of cardiac myxoma. No difference was found in the mortality between patients receiving an urgent and a delayed myxoma resection ($\chi^2=5.1$, $P=0.167$). Moreover, a staged cardiac myxoma resection was performed subsequent to other operations in 3 patients with an interval between operations of 1^[64], 2^[41] and 4^[80] weeks, respectively. Patients were followed up for 24.3±27.8 (range, 1-132; median, 12) months ($n=59$). Prognosis was reported in 72 patients: a full recovery in 41 (56.9%), an improvement in 13 (18.1%), unchanged in 2 (2.8%), cardiac myxoma recurrence in 5 (6.9%) and death in 11 (15.3%) patients.

By multivariate analysis, none of the independent variables including gender, age, comorbidities, multiplicity of stroke, peripheral embolic events, middle cerebral artery occlusion, basal ganglion infarct, left atrial myxoma, multiplicity of cardiac myxoma and surgical resection of cardiac myxoma was a predisposing risk factor for patient's mortality. Logistic regression analysis showed peripheral embolic

events ($P=0.024$) and non-surgical resection of cardiac myxoma ($P=0.033$) correlated significantly with mortality.

DISCUSSION

Cardiac myxoma is an important cause of stroke in young patients. Lee et al.^[46] reported that the stroke patients aged 48.5 (range, 17-70) years. Ekinici & Donnan's^[25] patient series aged between 6 and 82 years. The present study revealed young children at the age of 4 years can be a victim of cardiac myxoma stroke. Aziz et al.^[5] reported a female predominance with a female-male ratio of 2:1 in cardiac myxoma, and the present study also revealed a female predominance but with a smaller gender ratio. The classic symptom triad of cardiac myxoma patients includes obstructive, embolic and constitutional manifestations^[13]. Cardiac myxoma stroke is often of an acute onset, whereas tumor embolization including myxoma-induced cerebral aneurysm and myxomatous metastasis may show a delayed presentation^[46]. However, cardiac myxoma is responsible for only 0.5% of stroke, with females at the age of 50 years at greatest risk^[87]. Embolic manifestations occur in 20-45% of patients with a cardiac myxoma, sometimes as the onset symptom^[75], as a result of cerebral ischemia and less commonly hemorrhage^[18]. In a series of 113 atrial myxoma patients with neurologic presentations, 83% presented with ischemic stroke, most often at multiple sites (41%). Other manifestations included syncope (28%), psychiatric symptoms (23%), headache (15%) and seizures (12%)^[72]. Alvarez-Sabin et al.^[13] reported 11 of 28 (39.3%) cardiac myxoma patients had embolic phenomena, which affected the cerebral arteries in 6 (54.5%), peripheral arteries in 2 (18.2%) and both cerebral and peripheral arteries in 3 (27.3%) patients. Lee et al.^[45] reported 13 of 59 (22.0%) patients with a cardiac myxoma developed embolic events, 11 (18.6%) of which were in the brain, 2 (3.4%) in the limb and 1 (1.7%) in the eye. Furthermore, Yuan^[85] reported a patient with a delayed cardiac myxoma resection developed multiple embolic events in the lower extremities. Bajraktari et al.^[16] reported one cardiac myxoma patient who declined to all treatments for 7 years eventually developed mesenteric embolism.

The size of the atrial myxoma was variable with a mean of 2.7 (range, 0.4-6.5) cm as reported by Lee et al.^[46], and was 4.8±1.9 cm by Lee et al.^[45]. Cardiac myxoma was classified into two types according to their gross appearances: type 1, with an irregular or a villous surface and a soft consistency; and type 2, with a smooth surface and a compact consistency^[45]. Porapaktham et al.^[88] reported that cardiac myxomas with an irregular surface exceeded those with a smooth surface in number (62.2% vs. 37.8%). Embolic potential usually depends on the mobility other than on the size of the myxoma^[46]. Neurologic complications of atrial myxoma are most frequently cerebral infarcts caused by detached thrombus from the myxoma; while rarely by tumor fragments^[46].

Active illness is often accompanied by elevation of erythrocyte sedimentation rate and C-reactive protein, hyperglobulin-

Table 11. Surgical operations in 98 patients with cardiac myxoma stroke.

Surgical operation	n (%)
Isolated myxoma resection	82 (83.7)
Myxoma resection with concurrent heart operation	8 (8.2)
Mitral valve repair	3 (37.5)
Mitral valve replacement	1 (12.5)
Aortic valve replacement	1 (12.5)
Patent fossa ovalis/atrial septal defect closure	2 (25)
Coronary artery bypass grafting	1 (12.5)
Myxoma resection with other operation	6 (6.1)
Peripheral embolectomy	2 (33.3)
Peripheral embolectomy and amputation	1 (16.7)
Peripheral embolectomy, fasciotomies and amputation	1 (16.7)
Decompression craniotomy	1 (16.7)
Endovascular coiling of internal carotid artery aneurysm	1 (16.7)
No myxoma resection	2 (2.0)
Embolectomies and herniation decompressive hemicraniectomy	1 (50)
Cranial decompression	1 (50)

emia and anemia. Constitutional symptoms may be mediated by interleukin-6 produced by the myxoma itself^[89]. MRI is more sensitive than CT in identifying subtle abnormalities of the brain^[49]. Transthoracic and/or transesophageal echocardiography can be options for the screening of stroke of a cardiac source^[31].

The timing of treatment in such patient is always a matter of intense debate. Soleimanpour et al.^[74] proposed that the patients with a delayed presentation of more than 3-4.5 hours were not indicated for intravenous thrombolysis. Al-Said et al.^[11] disagreed the use of recombinant tissue plasminogen activator for intravenous thrombolysis due to probable myxoma source infarct. Vogel et al.^[80] suggested a 4-week delay of surgical resection of cardiac myxoma concerning the risk of intracerebral hemorrhage following cerebral infarction. Da Silva & de Freitas^[21] advocated a delayed surgery after a large stroke. However, Sethi^[90] carried out emergent peripheral vascular and cardiac surgeries in a patient with an acute large infarct of a cardiac myxoma origin and the patient did well.

Young adults, female predominance, single cerebral vessel (mostly the middle cerebral artery) and multiple territory involvements and solitary left atrial myxoma epitomize the outstanding characteristics of the patients with cardiac myxoma stroke. The most common affected cerebral vessel (the middle cerebral artery) and areas (the basal ganglion, cerebellum and parietal and temporal regions) correspond well to the common manifestations of conscious alteration, ataxia, hemiparesis and hemiplegia, aphasia and dysarthria. Initial CT scan carries a higher false negative rate in diagnosing cerebral infarction, and thus MRI should take the place of it, especially in patients with an immediate presentation. Echocardiography is a reliable means for determining the cardiogenic origin of stroke.

Limited patient information concerning the timing of onset, presentation and treatment as well as survival constitutes the main drawbacks of the present study. Multicenter prospective studies on this particular patient population are anticipated in the future.

CONCLUSIONS

Cardiac myxoma stroke is rare. Often does it affect young females. For an improved diagnostic accuracy, MRI and echocardiography are recommended for young stroke patients. Immediate thrombolytic therapy may completely resolve the cerebral stroke and improve the neurologic function of the patients. An early surgical resection of cardiac myxoma is recommended in patients with not large territory cerebral infarct so as to prevent from the potential consequences.

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Authors' roles & responsibilities

SMY	Main Author
GH	Final approval of the manuscript

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